## Amendment to the claims:

The current listing of claims as follows will replace all previously entered claims. :

## **Listing of Claims**

- 1. (Currently Amended) an-An Fc variant of a parent Fc polypeptide, said Fc variant comprising at least one amino acid modification in the Fc region of said parent Fc polypeptide said modification occurring at a position selected from the group consisting of: 228, 230, 231, 232, 240, 244, 245, 247, 262, 263, 266, 271, 273, 275, 281, 284, 291, 299, 302, 304, 313, 323, 325, 328, 332, and 336, wherein said Fc variant protein exhibits altered binding to an FcγR as compared to the parent Fc polypeptide, wherein numbering is according to the EU index.
- 2. (Previously Presented) An Fc variant according to claim 1, wherein said Fc variant comprises at least one substitution selected from the group consisting of: 230, 240, 244, 245, 247, 262, 263, 266, 273, 275, 299, 302, 313, 323, 325, 328, and 332.
- 3. (Previously Presented) An Fc variant according to claim 1, wherein said altered binding is an increase in affinity of said Fc variant to said FcyR.
- 4. (Previously Presented) An Fc variant according to claim 3, wherein said Fc variant binds with greater affinity to a mouse FcγR.
- 5. (Previously Presented) An Fc variant according to claim 3, wherein said said FcγR is a human Fc receptor selected from the group consisting of FcγRII, FcγRIIa, FcγRIIb, FcγRIIIa.
- 6. (Previously Presented) An Fc variant according to claim 3, wherein said Fc variant binds with greater affinity to human FcγRI and FcγRIIIa, but exhibits unaltered affinity to a human receptor selected from the group consisting of FcγRIIa, FcγRIIb, and FcγRIIc.
- 7. (Previously Presented) An Fc variant according to claim 6, wherein said Fc variant exhibits unaltered affinity to FcyRIIa, FcyRIIb, and FcyRIIc.
- 8. (Withdrawn) An Fc variant according to claim 3, wherein the affinity increase for binding to one or more human FcyRII's is greater than the affinity increase for binding to human FcyRII or FyRIIIa.
- (Withdrawn) An Fc variant according to claim 3, wherein the affinity increase is the same for binding to FcγRIIa, FcγRIIb, and FcγRIIc.

- 10. (Withdrawn) An Fc variant according to claim 3, wherein the affinity increase for binding to FcγRIIc is greater than the affinity increase for binding to FcγRIIb.
- 11. (Withdrawn) An Fc variant according to claim 3, wherein said Fc variant binds with greater affinity to human FcγRIIa, but exhibits unaltered affinity to a human receptor selected from the group consisting of FcγRI, FcγRIIb, FcγRIIc, and FcγRIIa.
- 12. (Withdrawn) An Fc variant according to claim 11, wherein said Fc variant exhibits unaltered affinity to FcyRI, FcyRIIb, FcyRIIc, and FcyRIIIa.
- 13. (Previously Presented) An Fc variant according to any of claims 1-12, wherein said Fc variant binds with greater affinity to FcRn.
- 14. (Withdrawn) An Fc variant according to any of claims 1-12, wherein said Fc variant binds with unaltered affinity to FcRn.
- 15. (Withdrawn) An Fc variant according to any of claims 1-12, wherein said Fc variant binds with reduced affinity to FcRn.
- 16. (Withdrawn Currently Amended) An Fc variant according to any of claims 1-12, wherein said Fc variant binds with greater affinity to C1q or enhances <u>complement dependent cytotoxicity (CDC)</u>.
- 17. (Previously Presented) An Fc variant according to any of claims 1-12, wherein said Fc variant binds with unaltered affinity to C1q or enhances CDC.
- 18. (Withdrawn) An Fc variant according to claim 1, wherein said Fc variant further comprises an engineered glycoform.
- 19. (Withdrawn) An Fc variant according to claim 18 wherein said engineered glycoform comprises an altered level of fucosylation or bisecting oligosaccharides as compared to the parent Fc polypeptide.
- 20. (Withdrawn) An Fc variant according to claim 18 wherein said engineered glycoform improves effector function.
- 21. (Currently Amended) An Fc variant according to claim 1, wherein said Fc variant modulates enhances or reduces an effector function selected from the group consisting of antibody

- dependent cell-mediated cytotoxicity (ADCC), CDC, or antibody dependent cell-mediated phagocytosis (ADCP) function as compared to the parent Fc polypeptide.
- 22. (Currently Amended) An Fc variant according to claim 21, wherein said effector function is antibody dependent cell-mediated cytotoxicity (ADCC).
- 23. (Previously Presented) An Fc variant according to claim 22, wherein said Fc variant improves ADCC in the presence of human effector cells as compared to said parent Fc polypeptide.
- 24. (Previously Presented) An Fc variant according to claim 22, wherein said ADCC improvement is an enhancement in potency such that the EC50 of said Fc variant is approximately 5-fold greater than that of said parent Fc polypeptide.
- 25. (Previously Presented) An Fc variant according to claim 23, wherein said ADCC improvement is an enhancement in potency such that the EC50 of said Fc variant is between approximately 5-fold and 1000-fold greater than that of said parent Fc polypeptide.
- 26. (Previously Presented) An Fc variant according to claim 23, wherein said ADCC improvement is an enhancement in efficacy such that the maximal ADCC is approximately 2-fold greater than that of said parent Fc polypeptide.
- 27. (Previously Presented) An Fc variant according to claim 26, wherein said Fc variant improves ADCC in the presence of mouse effector cells as compared to said parent Fc polypeptide.
- 28. (Previously Presented) An Fc variant according to claim 21, wherein said effector function is ADCP.
- 29. (Previously Presented) An Fc variant according to claim 28, wherein said Fc variant improves ADCP as compared to said parent Fc polypeptide.
- 30. (Previously Presented) An Fc variant according to claim 25 or 29, wherein CDC is unaffected.
- 31. (Previously Presented) An Fc variant according to claim 25 or 29, wherein CDC is ablated.
- 32. (Previously Presented) An Fc variant according to claim wherein said polypeptide has specificity for a target antigen selected from the group consisting of CD4, CD19, CD20, CD22, CD25, CD30,

CD33, CD52, CD80, B7-1, B7-2, CLTA-4, BAFF-R, Her2/neu, EGFR, EpCAM, MUC1, GD3, CEA, CA 125, HLA-DR, TNFalpha, and VEGF.

- 33. (Previously Presented) An Fc variant according to claim 1 or 21 comprising a modification selected from the group consisting of 228E, 228K, 228Y, 228G, 230E, 230Y, 230G, 231E, 231K, 231Y, 231P, 231G, 232E, 232K, 232Y, 232G, 262E, 262F, 271D, 271E, 271N, 271Q, 271K, 271R, 271S, 271T, 271H, 271A, 271V, 271L, 271I, 271F, 271M, 271Y, 271W, 271G, 275L, 281D, 281K, 281Y, 281P, 284E, 284N, 284T, 284L, 284Y, 291D, 291E, 291Q, 291T, 291H, 291I, 291G, 299D, 299E, 299N, 299Q, 299K, 299R, 299L, 299F, 299M, 299Y, 299W, 299P, 304D, 304N, 304T, 304H, 304L, 325K, 325R, 325S, 325F, 325M, 325Y, 325W, 325P, 325G, 328D, 328Q, 328K, 328R, 328S, 328T, 328V, 328I, 328Y, 328W, 328P, 328G, 332K, 332R, 332S, 332V, 332L, 332F, 332M, 332W, 332P, 332G, 336E, 336K, 336Y, 230A, 240A, 240I, 240M, 240T, 244H, 245A, 247G, 247V, 262A, 262E, 262I, 262T, 263A, 263I, 263M, 263T, 266A, 266I, 266M, 266T, 273I, 275W, 299A, 299E, 299F, 299H, 299I, 299L, 299S, 299V, 302I, 313F, 323I, 325A, 325D, 325E, 325H, 325I, 325L, 325Q, 325T, 325V, 328A, 328D, 328E, 328F, 328H, 328I, 328M, 328N, 328Q, 328T, 328V, 332A, 332D, 332E, 332H, 332N, 332Q, 332T, and 332Y.
- 34. (Previously Presented) An Fc variant according to claim 1 wherein said parent Fc polypeptide is an antibody comprising said Fc variant.
- 35. (Previously Presented) An Fc variant according to claim 1 wherein said Fc variant is an Fc fusion protein comprising said Fc variant.
  - 36. (Previously Presented) A pharmaceutical composition comprising an Fc variant according to claim 1 and a pharmaceutically acceptable carrier.
  - 37. (Withdrawn) A method of treating a mammal in need of said treatment, comprising administering a variant protein of claim 1.
  - 38. (Withdrawn) An Fc variant according to claims 1or 33 comprising at least one modification selected from the group consisting of A330L/I332E, D265F/N297E/I332E, D265Y/N297D/T299L/I332E, F241E/F243Q/V262T/V264E, F241E/F243Q/V262T/V264E/I332E,

F241E/F243R/V262E/V264R, F241E/F243R/V262E/V264R/I332E, F241E/F243Y/V262T/V264R. F241E/F243Y/V262T/V264R/I332E, F241L/F243L/V262I/V264I, F241L/V262I. F241R/F243Q/V262T/V264R, F241R/F243Q/V262T/V264R/I332E, F241W/F243W/V262A/V264A, F241Y/F243Y/V262T/V264T, F241Y/F243Y/V262T/V264T/N297D/I332E, F243L/V262I/V264W, F243L/V264I, L328D/I332E, L328E/I332E, L328H/I332E, L328I/I332E, L328M/I332E, L328N/I332E, L328N/I382E, L328N/I38P, L328N/I38P, L328N/I38P, L328N/I38P, L328N/I38P, L328N/I38P, L328N/I38P, L328N/I38P, L328Q/I332E, L328T/I332E, L328V/I332E, N297D/A330Y/I332E, N297D/I332E, N297D/I332E/S239D/A330L, N297D/S298A/A330Y/I332E, N297D/T299E/I332E, N297D/T299F/I332E, N297D/T299H/I332E, N297D/T299I/I332E, N297D/T299L/I332E, N297D/T299V/I332E, N297E/I332E, N297S/I332E, P230A/E233D/I332E, P244H/P245A/P247V, S239D/A330L/I332E, S239D/A330Y/I332E, S239D/A330Y/I332E/K326E, S239D/A330Y/I332E/K326T. S239D/A330Y/I332E/L234I, S239D/A330Y/I332E/L235D, S239D/A330Y/I332E/V240I, S239D/A330Y/I332E/V264T, S239D/A330Y/I332E/V266I, S239D/D265F/N297D/I332E, S239D/D265H/N297D/I332E, S239D/D265I/N297D/I332E, S239D/D265L/N297D/I332E, S239D/D265T/N297D/I332E, S239D/D265V/N297D/I332E, S239D/D265Y/N297D/I332E, S239D/I332D, S239D/I332E, S239D/I332E/A330I, S239D/I332N, S239D/I332Q, S239D/N297D/I332E. \$239D/N297D/I332E/A330Y, \$239D/N297D/I332E/A330Y/F241\$/F243H/V262T/V264T. S239D/N297D/I332E/K326E, S239D/N297D/I332E/L235D, S239D/S298A/I332E. S239D/V264I/A330L/I332E, S239D/V264I/I332E, S239D/V264I/S298A/I332E, S239E/D265N, S239E/D265Q, S239E/I332D, S239E/I332E, S239E/I332N, S239E/I332Q, S239E/N297D/I332E, S239E/V264I/A330Y/I332E, S239E/V264I/I332E, S239E/V264I/S298A/A330Y/I332E. S239N/A330L/I332E, S239N/A330Y/I332E, S239N/I332D, S239N/I332E, S239N/I332N, S239N/I332Q, S239N/S298A/I332E, S239Q/I332D, S239Q/I332E, S239Q/I332N, S239Q/I332Q, S239Q/V264I/I332E, S298A/I332E, V264E/N297D/I332E, V264I/A330L/I332E, V264I/A330Y/I332E, V264I/I332E, V264I/I33E, V264I/I3E, V26E, V264I/I3E, V26E, V264I/I3E, V26E, V264I/S298A/I332E, Y296D/N297D/I332E, Y296E/N297D/I332E, Y296H/N297D/I332E, Y296N/N297D/I332E, Y296Q/N297D/I332E, and Y296T/N297D/I332E.

39. (Withdrawn) An Fc variant according to claims 1, 33 or 38 further comprising a modification selected from the group consisting of 227E, 227K, 227Y, 227G, 233N, 233Q, 233K, 233R, 233S, 233T, 233H, 233A, 233V, 233L, 233I, 233F, 233M, 233Y, 233W, 233G, 234K, 234R, 234S, 234A, 234M, 234W, 234P, 234G, 235E, 235K, 235R, 235A, 235M, 235W, 235P, 235G, 236D, 236E, 236N, 236Q, 236K, 236R, 236S, 236T, 236H, 236A, 236V, 236L, 236I, 236F, 236M, 236Y, 236W, 236P, 237D, 237E, 237N, 237Q, 237K, 237R, 237S, 237T, 237H, 237V, 237L, 237I, 237F, 237M, 237Y, 237W, 237P, 238D, 238E, 238N, 238Q, 238K, 238R, 238S, 238T, 238H, 238V, 238L, 238I, 238F, 238M, 238Y, 238W, 238G, 239Q, 239K, 239P, 239L, 239I, 239M, 239W, 239P, 239G, 241D, 241E, 241Y, 243E, 246D, 246E, 246H, 246Y, 249Q, 249H, 249Y, 255E, 255Y, 258S, 258H, 258Y, 260D, 260E, 260H, 260Y, 264D, 264E, 264N, 264Q, 264K, 264R, 264S, 264H, 264W, 264P, 264G, 265Q, 265K, 265S, 265S, 265T, 265H, 265V, 265I, 265F, 265M, 265Y, 265W, 265P, 267E, 267Q, 267K,

267R, 267V, 267L, 267I, 267F, 267M, 267Y, 267W, 267P, 268D, 268E, 268Q, 268K, 268R, 268T, 268V, 268L, 268I, 268F, 268M, 268W, 268P, 268G, 269K, 269S, 269V, 269I, 269M, 269W, 269P, 269G, 270R, 270S, 270L, 270I, 270F, 270M, 270Y, 270W, 270P, 270G, 272D, 272R, 272T, 272H, 272V, 272L, 272F, 272M, 272W, 272P, 272G, 274D, 274N, 274S, 274H, 274V, 274I, 274F, 274M, 274W, 274P, 274G, 276D, 276T, 276H, 276V, 276I, 276F, 276M, 276W, 276P, 276G, 278D, 278N, 278Q, 278R, 278S, 278H, 278V, 278L, 278I, 278M, 278P, 278G, 280K, 280L, 280W, 280P, 280G, 282E, 282K, 282Y, 282P, 282G, 283K, 283H, 283L, 283Y, 283P, 283G, 285D, 285E, 285Q, 285K, 285Y, 285W, 286E, 286Y, 286P, 286G, 288D, 288E, 288Y, 290D, 290N, 290H, 290L, 290W, 292D, 292E, 292T, 292Y, 293N, 293R, 293S, 293T, 293H, 293V, 293L, 293I, 293F, 293M, 293Y, 293W, 293P, 293G, 294K, 294R, 294S, 294T, 294H, 294V, 294L, 294I, 294F, 294M, 294Y, 294W, 294P, 294G, 295D, 295E, 295N, 295R, 295S, 295T, 295H, 295V, 295I, 295F, 295M, 295Y, 295W, 295P, 295G, 296K, 296R, 296A, 296V, 296M, 296G, 297Q, 297K, 297R, 297T, 297H, 297V, 297L, 297I, 297F, 297M, 297Y, 297W, 297P, 297G, 298D, 298E, 298Q, 298K, 298R, 298I, 298F, 298M, 298Y, 298W, 300D, 300E, 300N, 300Q, 300K, 300R, 300S, 300T, 300H, 300A, 300V, 300M, 300W, 300P, 300G, 301D, 301E, 301H, 301Y, 303D, 303E, 303Y, 305E, 305T, 305Y, 317E, 317Q, 318Q, 318H, 318L, 318Y, 320N, 320S, 320H, 320V, 320L, 320F, 320Y, 320W, 320P, 320G, 322D, 322S, 322V, 322I, 322F, 322Y, 322W, 322P, 322G, 324H, 324F, 324M, 324W, 324P, 324G, 326P, 327E, 327K, 327R, 327H, 327V, 327I, 327F, 327M, 327Y, 327W, 327P, 329D, 329E, 329N, 329Q, 329K, 329R, 329S, 329T, 329H, 329V, 329L, 329I, 329M, 329Y, 329W, 329G, 330E, 330N, 330T, 330P, 330G, 331D, 331Q, 331R, 331T, 331L, 331I, 331F, 331M, 331Y, 331W, 333L, 333F, 333M, 333P, 334P, 335N, 335S, 335H, 335V, 335L, 335I, 335F, 335M, 335W, 335P, 335G, 337E, 337N, 337H, 233D, 234D, 234E, 234F, 234H, 234I, 234N, 234Q, 234T, 234V, 234Y, 235D, 235F, 235H, 235I, 235N, 235Q, 235S, 235T, 235V, 235Y, 239D, 239E, 239F, 239H, 239N, 239Q, 239T, 239Y, 241E, 241L, 241R, 241S, 241W, 241Y, 243H, 243L, 243Q, 243R, 243W, 243Y, 264A, 264E, 264F, 264I, 264L, 264M, 264R, 264T, 264W, 264Y, 265F, 265G, 265H, 265I, 265L, 265N, 265Q, 265T, 265V, 265Y, 267D, 267H, 267L, 267N, 267Q, 267T, 269F, 269H, 269L, 269N, 269R, 269T, 269Y, 270H, 270Q, 270T, 272I, 272K, 272S, 272Y, 274E, 274L, 274R, 274T, 274Y, 276E, 276L, 276R, 276S, 276Y, 278E, 278K, 278T. 278W, 283R, 296D, 296E, 296H, 296I, 296L, 296N, 296Q, 296S, 296T, 297D, 297E, 297S, 298A, 298H, 298N, 298T, 318R, 320D, 320I, 320T, 322H, 322T, 324D, 324I, 324L, 324R, 324T, 324V, 324Y, 326E, 326I, 326L, 326T, 327D, 327L, 327N, 327S, 327T, 329F, 330F, 330H, 330I, 330L, 330M, 330R, 330S, 330V, 330W, 330Y, 331H, 331V, 333A, 333H, 333I, 333T, 333Y, 334A, 334F, 334I, 334T, 335D, 335R, and 335Y.

40. (Withdrawn) An Fc variant according to claim 1 wherein said Fc variant further comprises a set of modifications selected from 241W/243W, 267Q/327S, 243L/264I, 234I/235D, 264E/297D/332E, 239E/265N, 239E/265Q, 239E/265G, 267L/327S and 240I/266I.

- 41. (Withdrawn) An Fc variant according to claim 33 wherein said antibody is a full length antibody.
- 42. (Withdrawn) An Fc variant according to claim 33 wherein said antibody is a human antibody.
- 43. (Withdrawn) An Fc variant according to claim 33 wherein said antibody is an antibody fragment.
- 44. (New) A Fc variant according to claim 1, wherein the Fc variant is an antibody.
- 45. (New) A Fc variant according to claim 1, wherein the Fc variant is an Fc fusion.